/ 9
Paper Number

REQUEST FOR ACCESS

	DATE: 4/26/94 SERIAL NUMBER: 42/98 599,543 FILING DATE: 10/18/90 APPLICANTS: Dependent
	FILING DATE: 10/18/90 APPLICANTS: Oppermann SIR: THE UNDERSIGNED HEREBY RESPECTFULLY REQUESTS ACCESS TO THE FILE
	HISTORY OF THE FOLLOWING ABANDONED APPLICATION WHICH WAS REFERRED TO IN U.S. PATENT NUMBER OR PRINTED APPLICATION
<i>!</i>	RESPECTFULLY SUBMITTED. The Land Bessenger
	OFFICIAL USE ONLY

Initials
File Information Unit

approved Mille - 4.26-94



United States Patent [19]

Oppermann et al.

[11] Patent Number:

5,266,683

[45] Date of Patent:

Nov. 30, 1993

[54] OSTEOGENIC PROTEINS

[75] Inventors: Hermann Oppermann, Medway;

Engin Ozkaynak, Milford; Thangavel Kuberasampath, Medway; David C. Rueger, Hopkinton; Roy H. L. Pang,

Medway, all of Mass.

[73] Assignee: Stryker Corporation, Kalamazoo,

Mich.

[*] Notice: The portion of the term of this patent

subsequent to Nov. 2, 2010 has been

disclaimed.

[21] Appl. No.: 841,646

[22] Filed: Feb. 21, 1992

Related U.S. Application Data

[60]	Continuation-in-part of Ser. No. 827,052, Jan. 28, 1992,
	Pat. No. 5,250,302, Ser. No. 579,865, Oct. 7, 1990, Pat.
	No. 5,108,753, Ser. No. 621,849, Dec. 4, 1990, aban-
	doned, Ser. No. 621,988, Dec. 4, 1990, abandoned, Ser.
	No. 810,560, Dec. 20, 1991, abandoned, Ser. No.
	569,920, Aug. 20, 1990, abandoned, Ser. No. 600,024,
	Oct. 18, 1990, abandoned, Ser. No. 599,543, Oct. 18,
	(1990, abandoned, Ser. No. 616,374, Nov. 21, 1990, Pat.
	No. 5,162,114, and Ser. No. 483,913, Feb. 22, 1990, Pat.
	No. 5,171,574, said Ser. No. 827,052, is a division of
	Ser. No. 179,406, Apr. 8, 1988, Pat. No. 4,568,590, said
	Ser. No. 579,865, is a division of Ser. No. 179,406, Apr.
	8, 1988, said Ser. No. 621,849, is a division of Ser. No.
	232,630, Aug. 15, 1988, abandoned, which is a con-
	tinuation-in-part of Ser. No. 179,406, Aug. 15, 1988,
	said Ser. No. 621,988, is a division of Ser. No. 315,342,
	Feb. 27, 1989, Pat. No. 5,011,691, which is a continua-
	tion-in-621t of Ser. No. 232,630, Feb. 23, 1989, said Ser.
	No. 810,560, is a continuation of Ser. No. 660,162, Feb.
	22, 1991, abandoned, which is a continuation of Ser.
	No. 422,699, Oct. 17, 1989, abandoned, which is a
	continuation-in-part of Ser. No. 315,342, Oct. 17, 1989,
	said Ser. No. 569,920, is a continuation-in-part of Ser.
	No. 422,699, Oci. 17, 1989, and Ser. No. 483,913, Oct.
	17, 1989, which is a continuation-in-part of Ser. No.
	472 612 Oct. 17 1000 Det No. 4 075 576 which is a
	422,613, Oct. 17, 1989, Pat. No. 4,975,526, which is a
	continuation-in-part of Ser. No. 315,342, Oct. 17, 1989,
	said Ser. No. 600,024, is a continuation-in-part of Ser.
	No. 569,920, Oct. 17, 1989, said Ser. No. 599,543, is a
	continuation-in-part of Ser. No. 569,920, Oct. 17, 1989.

[51]	Int. Cl. ⁵	A61K 37/U2; CU/K 5/W;
•		C07K 7/00; C07K 15/00
[52]	U.S. Cl	530/326; 530/327;
[]	530/32	28; 530/350; 530/395; 530/840
[58]	Field of Search	530/326, 327, 328, 395,
,		530/840, 300, 350

[56] References Cited U.S. PATENT DOCUMENTS

4.172.128	10/1979	Thiele et al 424/95
4.294.753	10/1981	Urie: 530/356
		Jefferies 424/15
		Seyedin et al 530/356
4.455.256	6/1984	Urist 530/356
4.563.350	1/1986	Nathan et al 424/95
		Urist 524/21
		continued on next page.)

FOREIGN PATENT DOCUMENTS

069260 6/1982 European Pat. Off. . (List continued on next page.)

OTHER PUBLICATIONS

Canalis et al., Science 210:1021-1023 (1980).
Glowacki et al., Lancet 4:959-963 (1981).
P. & di, Collagen Rel. Res. 1:209-226 (1981).
Sampath et al. Proc. Netl. Acad. Sci. USA 78:7599-7603 (1981).

(List continued on next page.)

Primary Examiner—Nathan M. Nutter Attorney, Agent, or Firm—Testa, Hurwitz & Thibeault

[57] ABSTRACT

Disclosed are (1) osteogenic devices comprising a matrix containing substantially pure natural-sourced mammalian osteogenic protein; (2) DNA and amino acid sequences for novel polypeptide chains useful as subunits of dimeric osteogenic proteins; (3) vectors carrying sequences encoding these novel polypeptide chains and host cells transfected with these vectors; (4) methods of producing these polypeptide chains using recombinant DNA technology; (5) antibodies specific for these novel polypeptide chains; (6) osteogenic devices comprising these recombinantly produced proteins in association with an appropriate carrier matrix; and (7) methods of using the osteogenic devices to mimic the natural course of endochondral bone formation in mammals.

58 Claims, 47 Drawing Sheets